

FIG. 1

Sequence of human APRIL (SEQ ID NOS: 1 and 2)

Human G70 cDNA (SEQ ID NO 1)

Length: 1465 bp

09855158.094

```
1  GCCAACCTTC CCTCCCCCAA CCCTGGGGCC GCCCCAGGGT TCCTGCGCAC
51  TGCCTGTTCC TCCTGGGTGT CACTGGCAGC CCTGTCCTTC CTAGAGGGAC
101 TGGAACCTAA TTCTCCTGAG GCTGAGGGAG GGTGGAGGGT CTCAAGGCAA
151 CGCTGGCCCC ACGACGGAGT GCCAGGAGCA CTAACAGTAC CCTTAGCTTG
201 CTTTCCTCCT CCCTCCTTTT TATTTTCAAG TTCCTTTTAA TTTCTCCTTG
251 CGTAACAACC TTCTTCCCTT CTGCACCACT GCCCGTACCC TTACCCGCCC
301 CGCCACCTCC TTGCTACCCC ACTCTTGAAA CCACAGCTGT TGGCAGGGTC
351 CCCAGCTCAT GCCAGCCTCA TCTCCTTTCT TGCTAGCCCC CAAAGGGCCT
401 CCAGGCAACA TGGGGGGCCC AGTCAGAGAG CCGGCACTCT CAGTTGCCCT
451 CTGGTTGAGT TGGGGGGCAG CTCTGGGGGC CGTGGCTTGT GCCATGGCTC
501 TGCTGACCCA ACAAACAGAG CTGCAGAGCC TCAGGAGAGA GGTGAGCCGG
551 CTGCAGGGGA CAGGAGGCCC CTCCCAGAAT GGGGAAGGGT ATCCCTGGCA
601 GAGTCTCCCG GAGCAGAGTT CCGATGCCCT GGAAGCCTGG GAGAGTGGGG
651 AGAGATCCCG GAAAAGGAGA GCAGTGCTCA CCCAAAAACA GAAGAAGCAG
701 CACTCTGTCC TGCACCTGGT TCCCATTAAAC GCCACCTCCA AGGATGACTC
751 CGATGTGACA GAGGTGATGT GGCAACCAGC TCTTAGGCGT GGGAGAGGCC
801 TACAGGCCCA AGGATATGGT GTCCGAATCC AGGATGCTGG AGTTTATCTG
851 CTGTATAGCC AGGTCTGTGT TCAAGACGTG ACTTTCACCA TGGGTCAGGT
901 GGTGTCTCGA GAAGGCCAAG GAAGGCAGGA GACTCTATTC CGATGTATAA
951 GAAGTATGCC CTCCCACCCG GACCGGGCCT ACAACAGCTG CTATAGCGCA
1001 GGTGTCTTCC ATTTACACCA AGGGGATATT CTGAGTGTCA TAATTCCCCG
1051 GGCAAGGGCG AAACCTTAACC TCTCTCCACA TGGAACCTTC CTGGGGTTTG
1101 TGAAACTGTG ATTGTGTTAT AAAAAGTGGC TCCCAGCTTG GAAGACCAGG
1151 GTGGGTACAT ACTGGAGACA GCCAAGAGCT GAGTATATAA AGGAGAGGGA
1201 ATGTGCAGGA ACAGAGGCGT CTTCTGGGT TTGGCTCCCC GTTCCTCACT
1251 TTTCCCTTTT CATTCCCACC CCCTAGACTT TGATTTTACG GATATCTTGC
1301 TTCTGTTCCC CATGGAGCTC CGAATTCTTG CGTGTGTGTA GATGAGGGGC
1351 GGGGACGGG CGCCAGGCAT TGTCAGACC TGGTCGGGGC CCACTGGAAG
1401 CATCCAGAAC AGCACCACCA TCTAACGGCC GCTCGAGGGA AGCACCCGGC
1451 GGTGTTGGCG AAGTC
```

The proposed transmembrane domains are boxed

human G70 protein sequence (SEQ ID NO 2)

```
1  MPASSPFLLA PKGPPGNMGG PVREPALSVA LWLSWGAALG AVACAMALLT
51  QQTELOSLRR EVSRLQGTGG PSQNGEGYPW QSLPEQSSDA LEAWESGERS
101 RKRRAVLTQK QKKQHSVLHL VPINATSKDD SDVTEVMWQP ALRRGRGLQA
151 QGYGVRIQDA GYLLYSQVL FQDVTFTMGQ VVSREGQGRQ ETLFRCIRSM
201 PSHPDRAVNS CYSAGVFHLH QGDILSVIIP RARAKLNLSP HGTFLGFV
```

FIG. 2A

Sequence of mouse G70 (SEQ ID NOS: 3 and 4)

Mouse G70 (SEQ ID NO 3)

1 CATGCCGAGT GCTTTGTGTG TGTTACCTGC TCTAAGAAGC TGGCTGGGCA
51 GCGTTTCACC GCTGTGGAGG ACCAGTATTA CTGCGTGGAT TGCTACAAGA
101 ACTTTGTGGC CAAGAAGTGT GCTGGATGCA AGAACCCCAT CACTGGGTTT
151 GGTAAAGGCT CCAGTGTGGT GGCCTATGAA GGACAATCCT GGCACGACTA
201 CTGCTTCCAC TGCAAAAAAT GCTCCGTGAA TCTGGCCAAC AAGCGCTTTG
251 TATTTCATAA TGAGCAGGTG TATTGCCCTG ACTGTGCCAA AAAGCTGTAA
301 CTTGACGGCT GCCCTGTCCT TCCTAGATAA TGGCACCAA TTCTCCTGAG
351 GCTAGGGGGG AAGGAGTGTC AGAGTGTAC TAGCTCGACC CTGGGGACAA
401 GGGGGACTAA TAGTACCCTA GCTTGATTTC TTCCTATTCT CAAGTTCCTT
451 TTTATTTCTC CCTTGCGTAA CCCGCTCTTC CTTCTGTGC CTTTGCCTGT
501 ATTCCCACCC TCCCTGCTAC CTCTTGCCA CCTCACTTCT GAGACCACAG
551 CTGTTGGCAG GGTCCCTAGC TCATGCCAGC CTCATCTCCA GGCCACATGG
601 GGGGCTCAGT CAGAGAGCCA GCCCTTTCGG TTGCTCTTTG GTTGAGTTGG
651 GGGGCAGTTC TGGGGGCTGT GACTTGTGCT GTCGCACTAC TGATCCAACA
701 GACAGAGCTG CAAAGCCTAA GGCGGGAGGT GAGCCGGCTG CAGCGGAGTG
751 GAGGGCCTTC CCAGAAGCAG GGAGAGCGCC CATGGCAGAG CCTCTGGGAG
801 CAGAGTCCTG ATGTCCTGGA AGCCTGGAAG GATGGGGCGA AATCTCGGAG
851 AAGGAGAGCA GTACTCACCC AGAAGCACAA GAAGAAGCAC TCAGTCCTGC
901 ATCTTGTTCC AGTTAACATT ACCTCCAAGG ACTCTGACGT GACAGAGGTG
951 ATGTGGCAAC CAGTACTTAG GCGTGGGAGA GGCCTGGAGG CCCAGGGAGA
1001 CATTGTACGA GTCTGGGACA CTGGAATTTA TCTGCTCTAT AGTCAGGTCC
1051 TGTTTCATGA TGTGACTTTC ACAATGGGTC AGGTGGTATC TCGGGAAGGA
1101 CAAGGGAGAA GAGAACTCT ATTCCGATGT ATCAGAAGTA TGCCTTCTGA
1151 TCCTGACCGT GCCTACAATA GCTGCTACAG TGCAGGTGTC TTTCATTTAC
1201 ATCAAGGGGA TATTATCACT GTCAAAATTC CACGGGCAAA CGCAAACTT
1251 AGCCTTTCTC CGCATGGAAC ATTCTGGGG TTTGTGAAAC TATGATTGTT
1301 ATAAAGGGGG TGGGGATTTC CCATTCCAAA AACTGGCTAG ACAAAGGACA
1351 AGGAACGGTC AAGAACAGCT CTCCATGGCT TTGCCTTGAC TGTTGTTCCCT
1401 CCCTTTGCCT TTCCCCTCC CACTATCTGG GCTTTGACTC CATGGATATT
1451 AAAAAAGTAG AATATTTTGT GTTTATCTCC CAAAAA

Sequence ID: 3



FIG. 2B

Mouse G70 Length: 241 (SEQ ID NO 4)

1 MPASSPGHMG GSVREPALSV ALWLSWGAVL GAVTCAVALL IQQTELQSLR
51 REVSRLQMSG GPSQKQGERP WQSLWEQSPD VLEAWKDGAK SRRRRAVLTQ
101 KHKKKHSVLH LVPVNITSKD SDVTEVMWQP VLRRGRGLEA QGDIVRVWDT
151 GIYLLYSQVL FHDVTFTMGQ VVSREGQGRR ETLFRCIRSM PSDPDRAVNS
201 CYSAGVFHLH QGDIITVKIP RANAKLSLSP HGTFLGFVKL *

G-70 FLAG des92 (smuG70) Strain #4081 (SEQ ID NO 19):

MDYKDDDDKKKKKHSVLH LVPVNITSKDS DVTEVMWQPYLRRGRGLEA QGDIVRVWDTGIY
LLYSQVLFHDVTFTMGQ VVSREGQGRRET LFCIRSMPSDPDRAYNSCYSAGVFHLHQGDII
TVKIPRANAKLSLSPHGTFLGFVKL*

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FIG. 3

FIG. 3

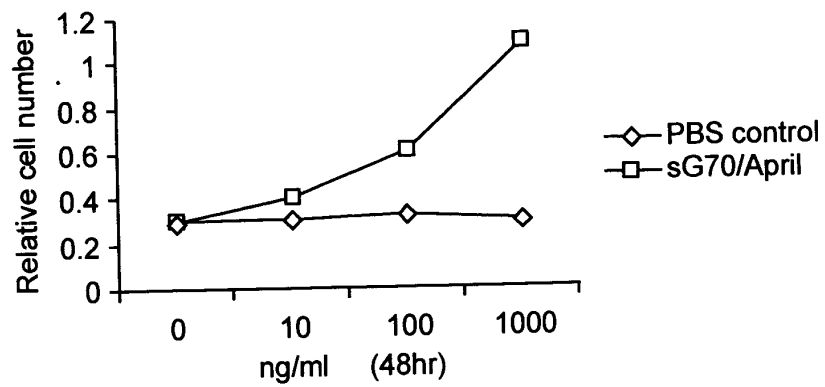
Alignm. of human and mouse G70

| | | | | | | |
|--------|-----|--|----------------------------------|------------------|-------------|-----------|
| mouse: | 1 | MPASS-----PGHMGGS | VREPALSVALWLSWGA | VLGAVTCAVALL | IQQTELSLRR | 51 |
| | | MPASS | PG+MGG | VREPALSVALWLSWGA | LGAV CA+ALL | QQTELSLRR |
| human: | 1 | MPASSPFLAPKPPGNMGGP | VREPALSVALWLSWGA | ALGAVACAMALL | TQQTELSLRR | 60 |
| mouse: | 52 | EVSRLQSGGPSQKQGERPWQSLWEQSPDVLEAWKDGAKSRRRRAVLTQKHKKKHVS | LHL | | | 111 |
| | | EVSRLQ +GGPSQ | PWQSL EQS D LEAW+ G +SR+RRAVLTQK | KK+HVS | LHL | 120 |
| human: | 61 | EVSRLQGTGGPSQNGEGYPWQSLPEQSSDALEAWESGERSRKRRAVLTQKQKKQHS | VHL | | | 170 |
| mouse: | 112 | VPVNITSKD-SDVTEVMWQPVLRGRGRGLEAQQDIVRVWDTGIYLLYSQVLFHDVFTMGQ | | | | 180 |
| | | VP+N TSKD SDVTEVMWQP LRRGRGL+AQG | VR+ D G+YLLYSQVLF | DVFTMGQ | | 230 |
| human: | 121 | VPINATSKDDSDVTEVMWQPALRRGRGLQAQGYGVRIQDAGVYLLYSQVLFQDVFTMGQ | | | | 240 |
| mouse: | 171 | VVSREGQRRRETLFRCIRSMPSDPDRAYNSCYSAGVFHLHQGDIITVKIPRANAKLSLSP | | | | 230 |
| | | VVSREGQGR+ETLFRCIRSMPS | PDRAYNSCYSAGVFHLHQGDI++V | IPRA AKL+LSP | | 240 |
| human: | 181 | VVSREGQGRQETLFRCIRSMPSHPDRAYNSCYSAGVFHLHQGDILSVIIPRARKLNLS | P | | | 240 |
| mouse: | 231 | HGTFLGFVKL | 240 | | | |
| | | HGTFLGFVKL | | | | |
| human: | 241 | HGTFLGFVKL | 250 | | | |

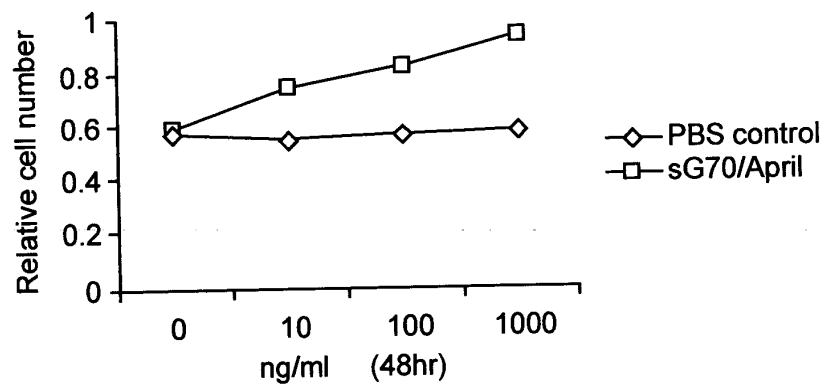


FIG. 4A

Effect of sG70/April on Raji cell proliferation



Effect of sG70/April on Jurkat cell proliferation



Effect of sG70/April on K562 cell proliferation

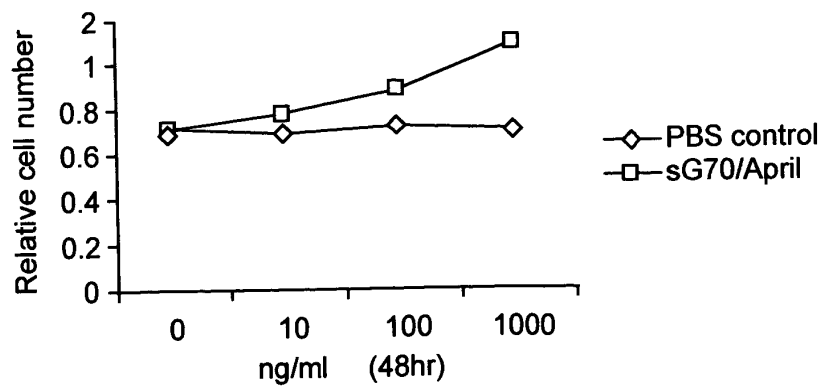
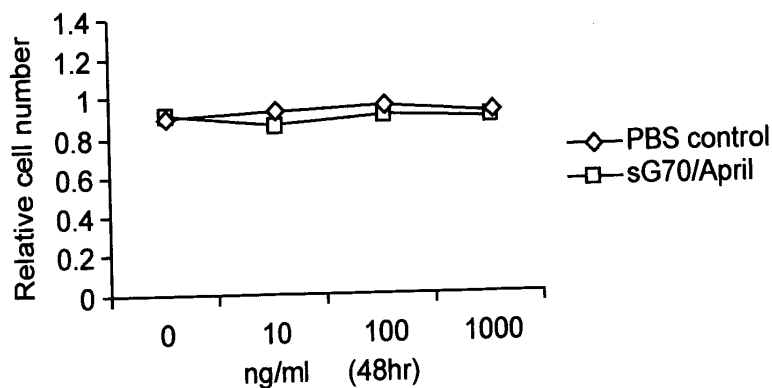


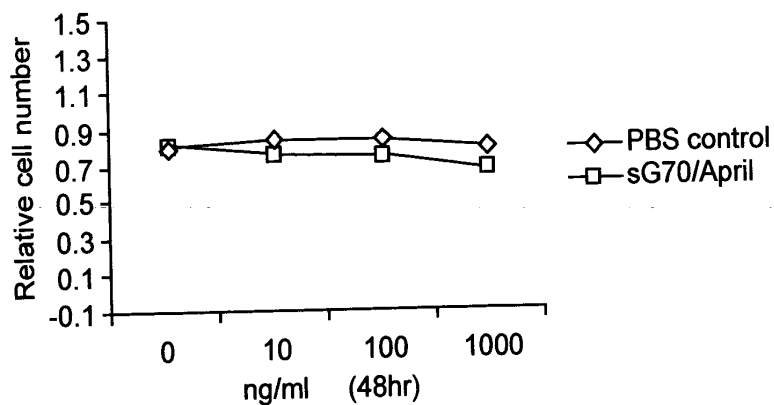


FIG. 4B

Effect of sG70/April on U937 cell proliferation



Effect of sG70/April on 293 T cell proliferation



Effect of sG70/April on 3T3 cell proliferation

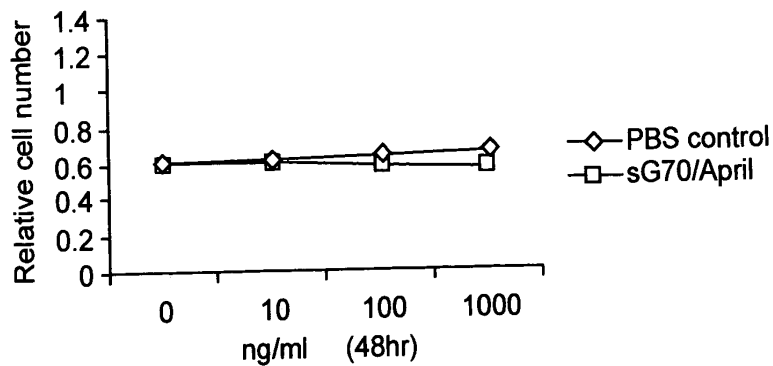


FIG. 5A

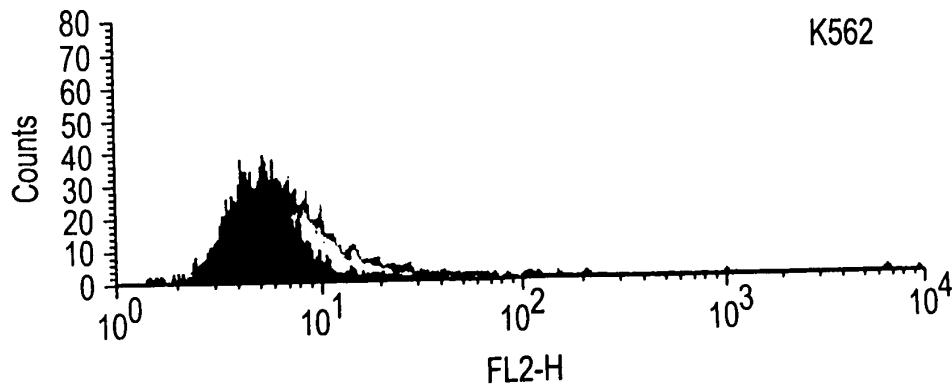
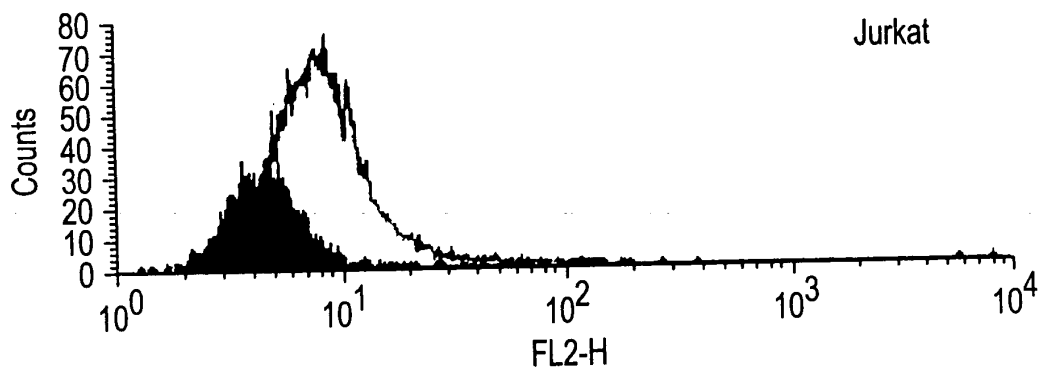
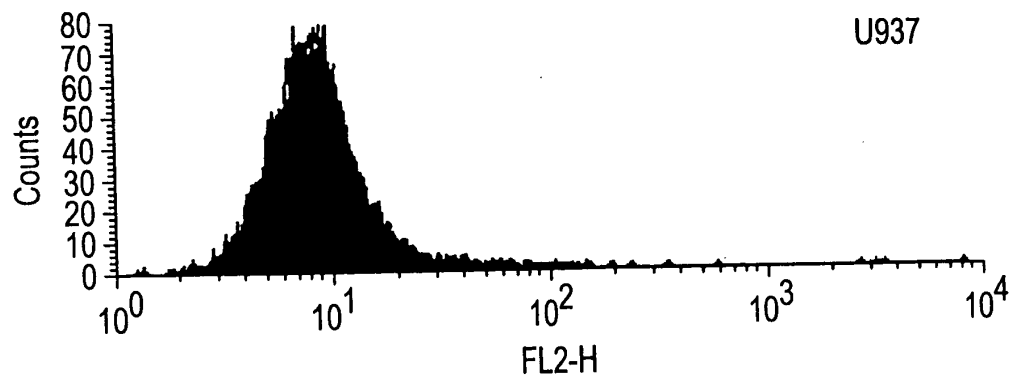




FIG. 5B-1

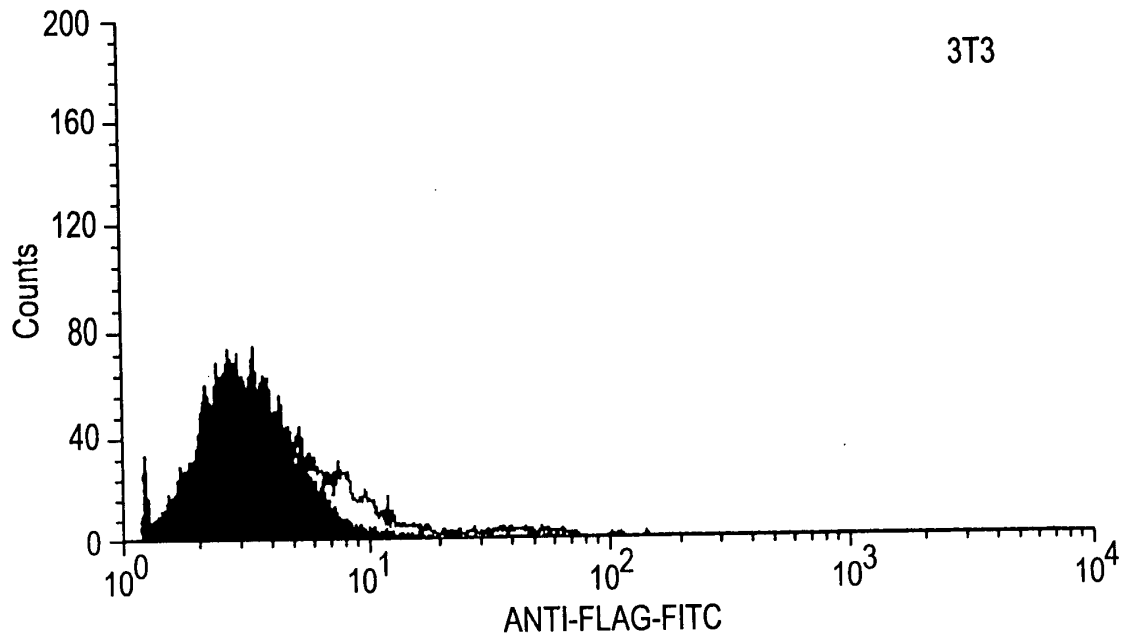


FIG. 5B-2

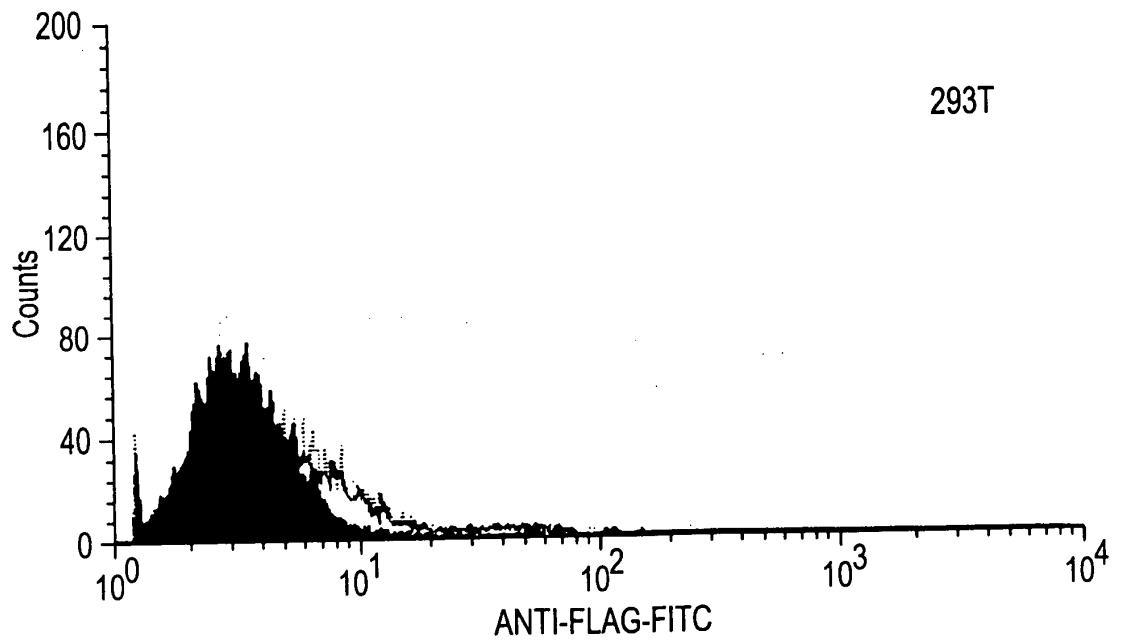




FIG. 5B-3

FIG. 5B-3

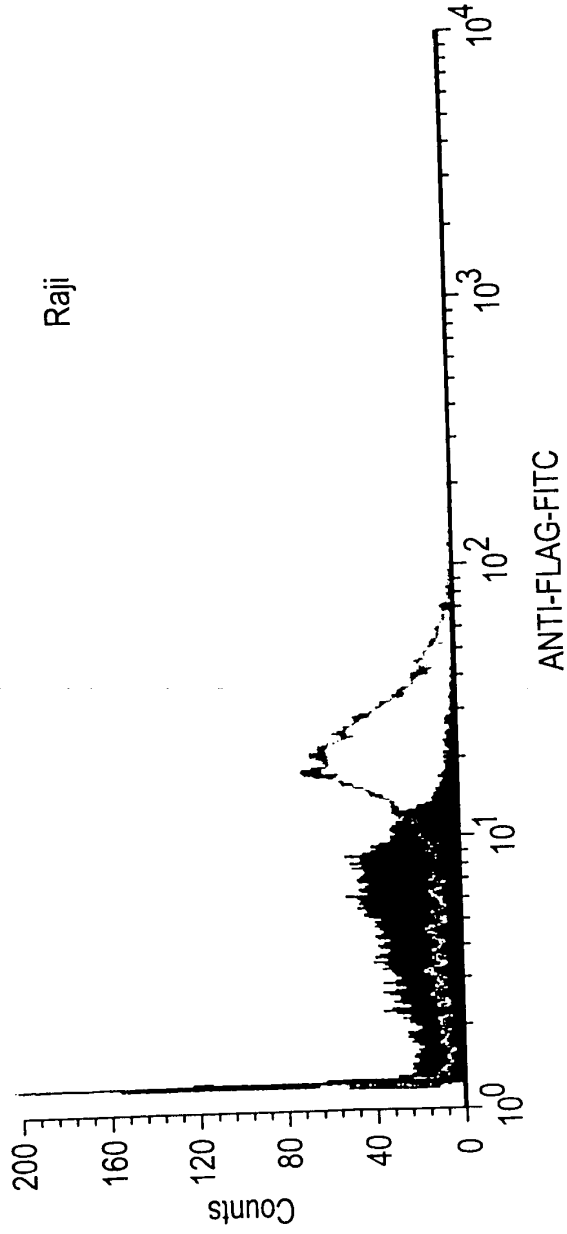
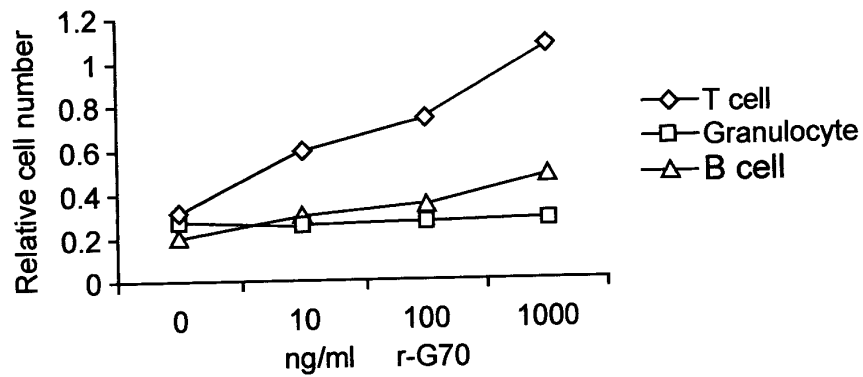




FIG. 6

The effect of r-G70/April on human
peripheral blood B cell, T cell and Granulocyte



The effect of IL-2 and G70/April on human
peripheral T cell proliferation

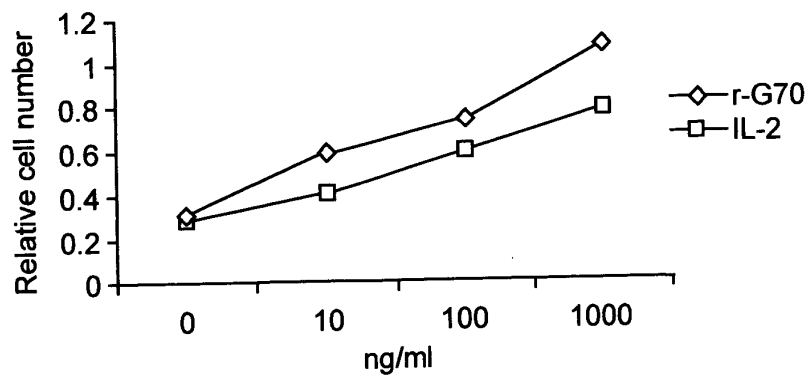
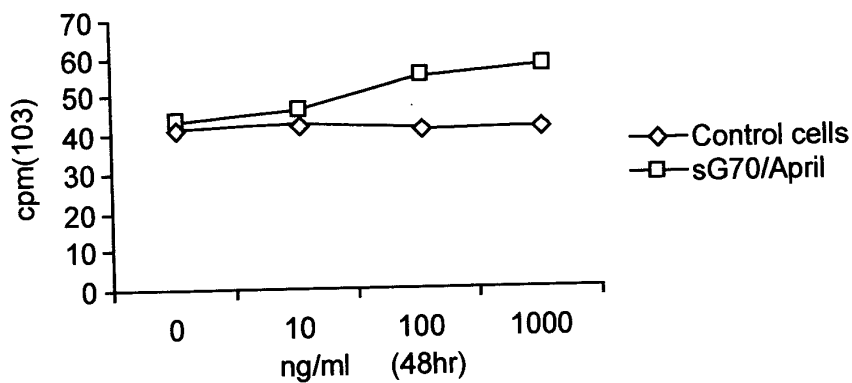




FIG. 7

Effect of sG70/April on murine B cell proliferation



Effect of sG70/April on murine T cell proliferation

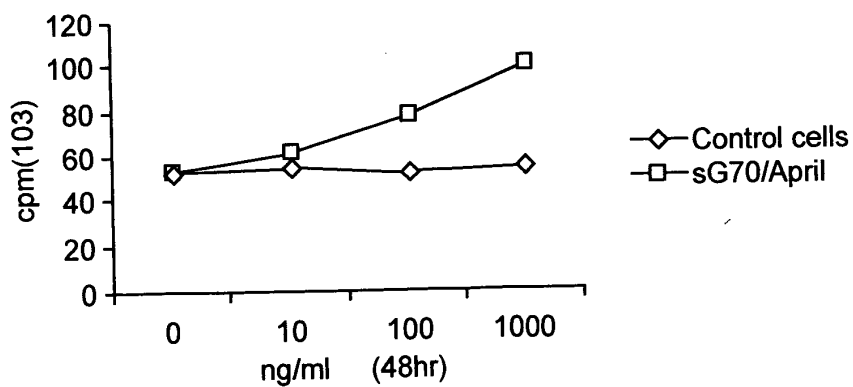




FIG. 8

Effect of G70/April on murine T cell
proliferation costimulated through CD28
antibody

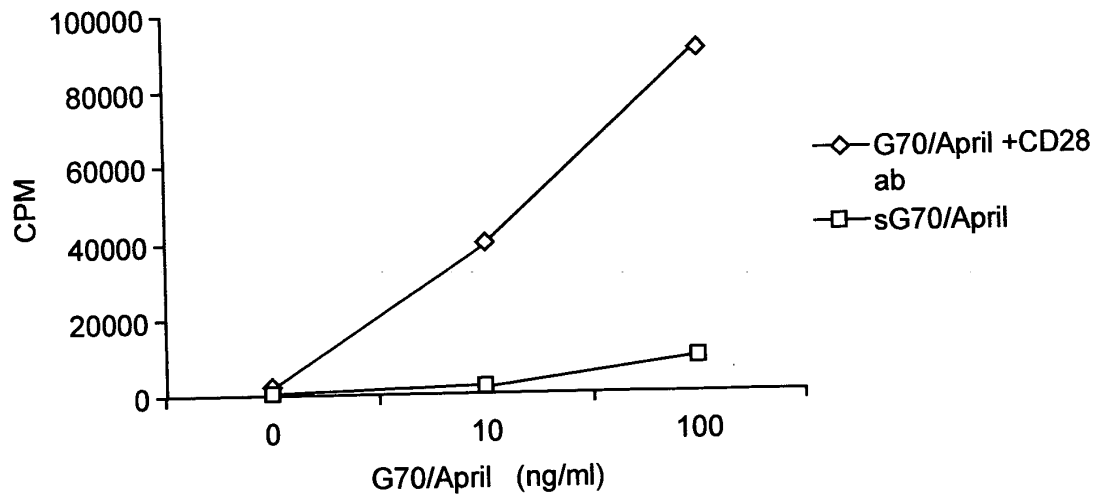




FIG. 9

Co-stimulatory activity of G70/April on mouse T cells

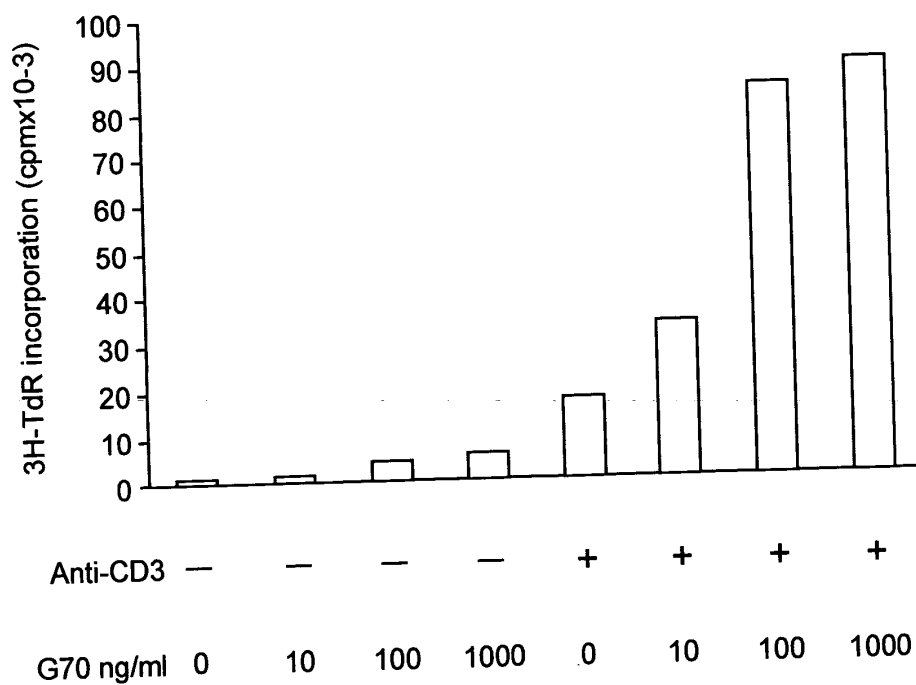




FIG. 10A

Human BCMA

Human (SEQ ID NO: 5):

1 MAGQCSQNEY FDSLLHACIP CQLRCSSNTP PLTCQRYCNA SVTNSVKGTN
51 AILWTCLGLS LIISLAVFVL MFLLRKISSE PLKDEFKNTG SGLLGMANID
101 LEKSRTGDEI ILPRGLEYTV EECTCEDCIK SKPKVSDHC FPLPAMEEGA
151 TILVTTKTND YCKSLPAALS ATEIEKSISA R

Human (SEQ ID NO: 5):

MAGQCSQ⁷ ⁸NEYFDSLLHA CIPCQLRCSS NTPPLTCQRY CNASVTNSVK
GTNA ILWTCL GLSLIISLAV FVLMFLLRKI SSEPLKDEFK NTGSGLLGMA
NIDLEKSRTG DEIILPRGLE YTVEECTCED CIKSKPKVDS DHCFLPAME
EGATILVTTK TNDYCKSLPA ALSATEIEKS ISAR

hBCMA's extracellular domain (SEQ ID NO: 6):

MAGQCSQ (NEYFDSLLHA CIPCQLRCSS NTPPLTCQRY)CNASVTNSVK
GTNA

hBCMA's cysteine-rich consensus region (SEQ ID NO: 7):

CSQ NEYFDSLLHA CIPCQLRCSS NTPPLTCQRY C

hBCMA's transmembrane region (SEQ ID NO: 8):

ILWTCL GLSLIISLAV FVLMF

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FIG. 10B

huBCMA-Fc (SEQ ID NO: 9):

MAGQCSQNEYFDSLLHACIPCQLRCSSNTPPLTCQRYCNASVTNSVKGTNAGGG
GGDKTHTCPPCPAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVDVSHEDPEVK
FNWYVDGVEVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKAL
PAPIEKTISKAKGQPREPQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWESNG
QPENNYKTTTPVLDSDGSFFLYSKLTVDKSRWQQGNVSCSVMHEALHNHYTQKS
LSLSPGK*

muBCMA-Fc (SEQ ID NO: 10):

MAQQCFHSEYFDSLLHACKPCHLRCSNPPATCQPYCDPSVTSSVKGSYTGGGGG
DKTHTCPPCPAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVDVSHEDPEVKFN
WYVDGVEVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPA
PIEKTISKAKGQPREPQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWESNGQP
ENNYKTTTPVLDSDGSFFLYSKLTVDKSRWQQGNVSCSVMHEALHNHYTQKSLS
LSPGK*

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FIG. 11

FIG. 11

Alignment of human BCMA amino acid sequence and murine BCMA amino acid sequence

murine BCMA amino acid sequence Length: 185 (SEQ ID NO: 11):

1 MAQQCFHSEY FDSLLHACKP CHLRCSNPPA TCQPYCDPSV TSSVKGTYTV
51 LWIFLGLTLV LSLALFTISF LLRKMNPEAL KDEPQSPGQL DGSQAQDKAD
101 TELTRIRAGD DRIFPRSLEY TVEECTCEDC VKSKPKGDS D HFFPLPAMEE
151 GATILVTTKT GDYKSSVPT ALQSVGMGEK PTHTR

alignment of human BCMA amino acid sequence and murine BCMA amino acid sequence.

Query: 4 MAGQCSQNE~~Y~~FDSLLHACIP~~C~~QLRCS~~S~~NT~~P~~PLTCQRYCNASVTNSVKGTNAILWTCGLGLS 63
MA QC ~~Y~~FDSLLHAC PC LRCS~~+~~ PP TCQ YC+ SVT+SVKGT +LW LGL+
Sbjct: 1 MAQQCFHSEYFDSLLHACKPCHLRCSN--PPATCQPYCDPSVTSSVKGTYTVLWIFLGLT 58
64 LIISLAVFVLMFLLRKISSEPLKDEFKNTG----SGLLGMANIDLEKSRGTGEIILPRGL 119
L++SLA+F + FLLRK++ E LKDE ++ G S L A+ +L + R GD+ I PR L
Sbjct: 59 LVLSLALFTISFLLRKMNPEALKDEPQSPGQLDGSQAQDKADTELTRIRAGDDRIFFRSL 118
120 EYTVEECTCEDCIKSKPKVSDSDHCFFLPAMEEGATILVTTKTNDYCKS-LPAAL-SATEI 177
EYTVEECTCEDC+KSKPK DSDH FPLPAMEEGATILVTTKT DY KS +P AL S +
Sbjct: 119 EYTVEECTCEDCVKSKPKGSDSDHFFPLPAMEEGATILVTTKTGDYKSSVPTALQSVGM 178
Query: 178 EKSISAR 184
EK R
Sbjct: 179 EKPTHTR 185

Human TACI

[illegible]

MSGLGRSRRGGRSRVDQEERFPQGLWTGVAMRSCPEEQYWDPLLGTCMSC
KTICNHQSQRTCAAFCRSLSCRKEQGKEYDHLRLDCISCASICGQHPKQC
AYFCENKLRSPVNLPELRRQRSGEVENNSDNSGRYQGLEHRGSEASPAL
PGLKLSADQVALVYSTLGLCLCAVLCCFLVAVACFLKKRGDPCSCQPRSR
PRQSPA KSSQDHAMEAGSPVSTSPEPVETCSFCFPECRAPTQESAVTPGT
PDPTCAGRWGCHTRTTVLQPCPHIPDSGLGIVCVPAQEGGPGA

1 MSGLGSRRRG GRSRVDQEER FPQGLWTGVA MRSCPEEQYW DPLLGTCSMC
51 KTICNHQSQR TCAAFCSLS CRKEQGKFYD HLLRDCISCA SICGQHPKQC
101 AYFCENKLRS PVNLPPELRR QRSGEVENNS DNSGRYQGLE HRGSEASPAL
151 PGLKLSADQV ALVYST



FIG. 12B

huTACI's cysteine-rich consensus region (SEQ ID NO: 16):
CPPEQYWDPLLGTCSCKTICNHQSQR TCAAF C and
CRKEQGKFYDHLRDCISCASICGQHPKQ CAYFC

transmembrane region (SEQ ID NO: 17):
LGLCLCAVLCCFLVAVACFL

hTACI-Fc (SEQ ID NO: 18):

1 MSGLGRSRRG GRSRVDQEER FPQGLWTGVA MRSCPEEQYW DPLLGTCSMC
51 KTICNHQSQR TCAAFCRSL S CRKEQGKFYD HLLRDCISCA SICGQHPKQC
101 AYFCENKLRS PVNLPPELRR QRSGEVENNS DNSGRYQGLE HRGSEASPAL
151 PGLKLSADQV ALVYSGGGGG DKHTTCPPCP APELLGGPSV FLFPPKPKDT
201 LMISRTPEVT CVVVDVSHED PEVKFNWYVD GVEVHNAKTK PREEQYNSTY
251 RVSVLTVLH QDWLNGKEYK CKVSNKALPA PIEKTISKAK GQPREPQVYT
301 LPPSRDELTK NQVSLTCLVK GFYPSDIAVE WESNGQPENN YKTTTPVLDS
351 DGSFFLYSKL TVDKSRWQQG NVFSCSV MHE ALHNHYTQKS LSLSPGK*

```

34 CPEEQYWDPLLGTCSCKTICNHQS.QRTCAAFCSRSLSCRKEQGKFYDHL 82
   | : : | . | | . | . | . | | : | . | . :
8  CSQNEYFDSLHACIPCQLRCSNTPPLTCQRYCNASVTNSVKGT..NAI 55
      .
      83 LRDCISCASI 92
      | | : . |
      56 LWTCLGLSLI 65

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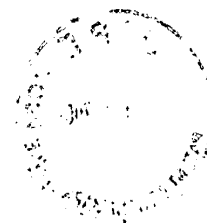


FIG. 14A

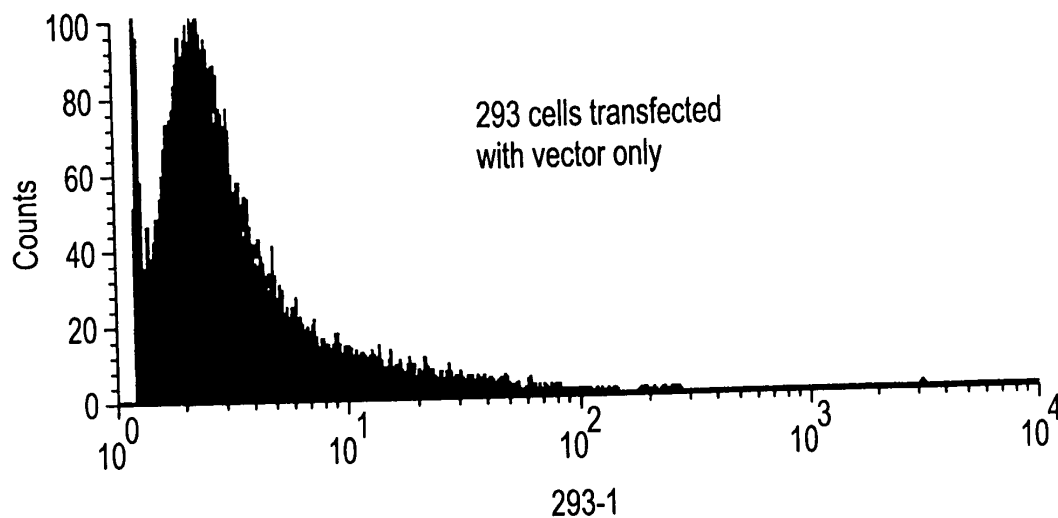


FIG. 14B

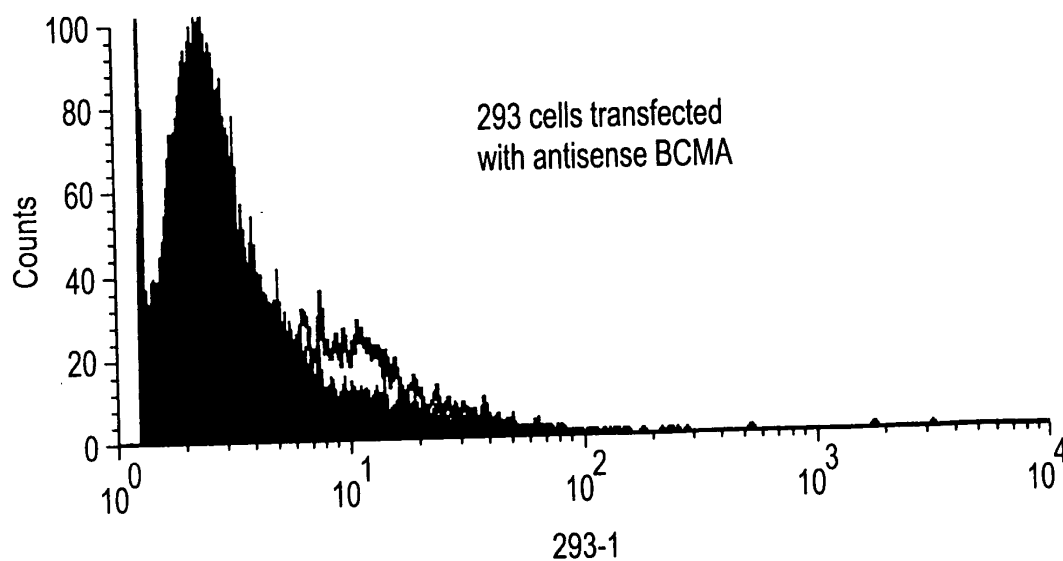


FIG. 14A



FIG. 14C

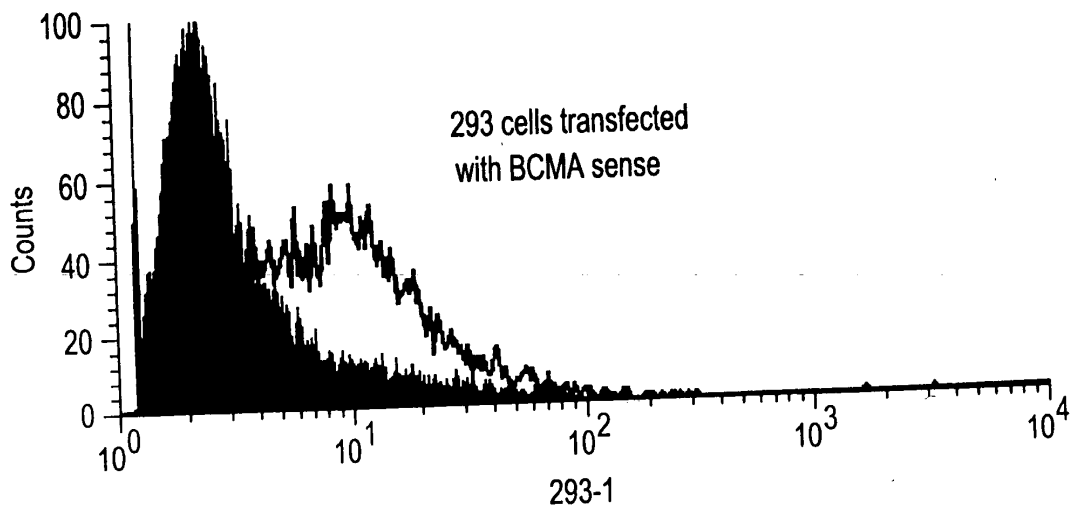




FIG. 15A

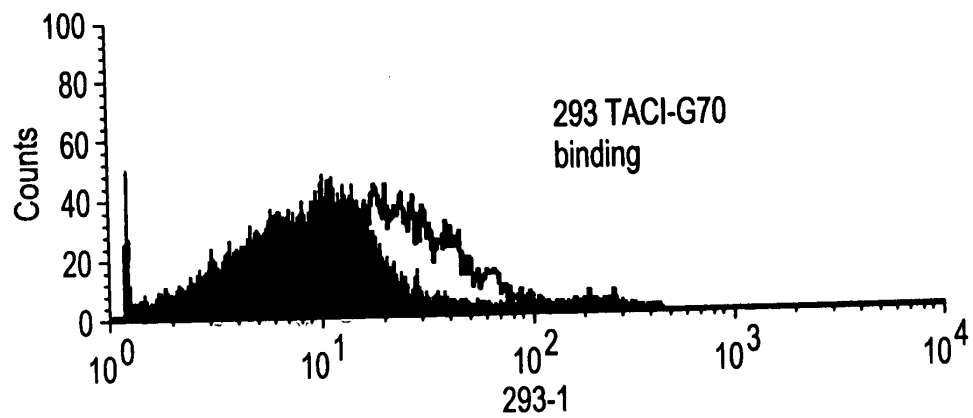


FIG. 15B

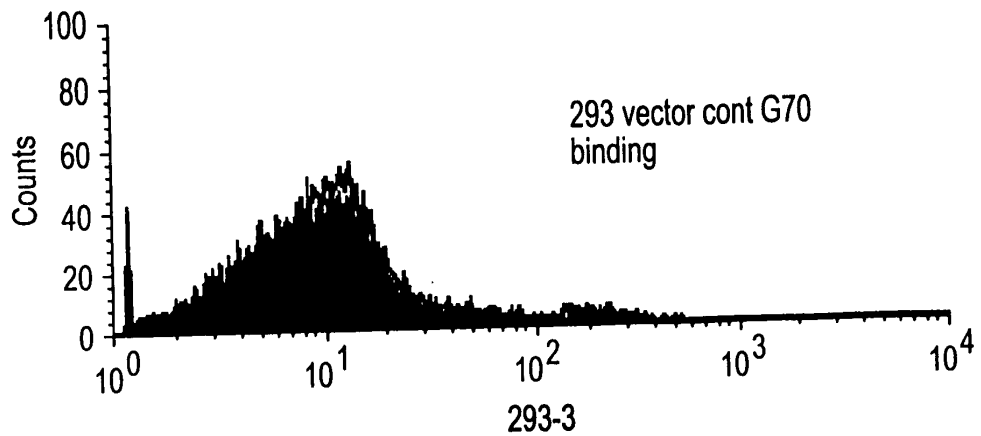


FIG. 16A

FIG. 16A

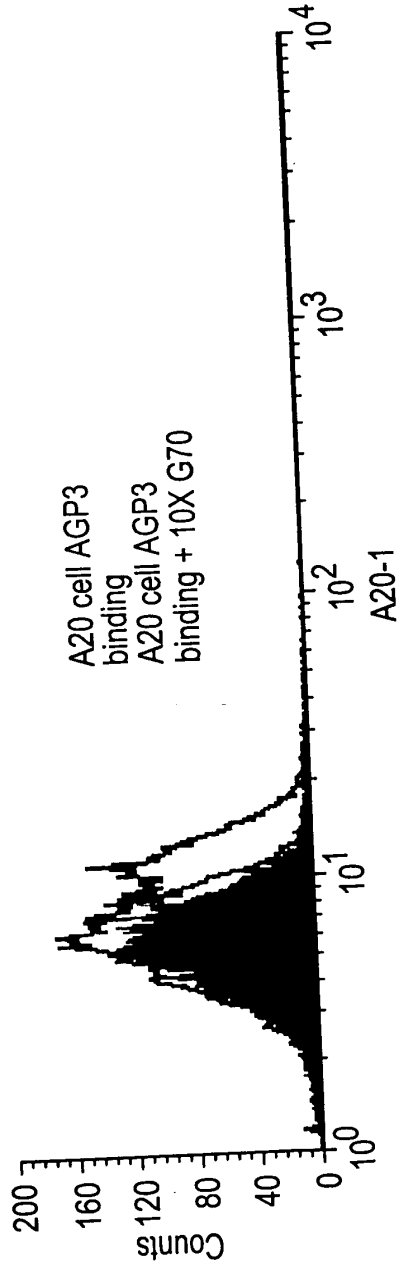
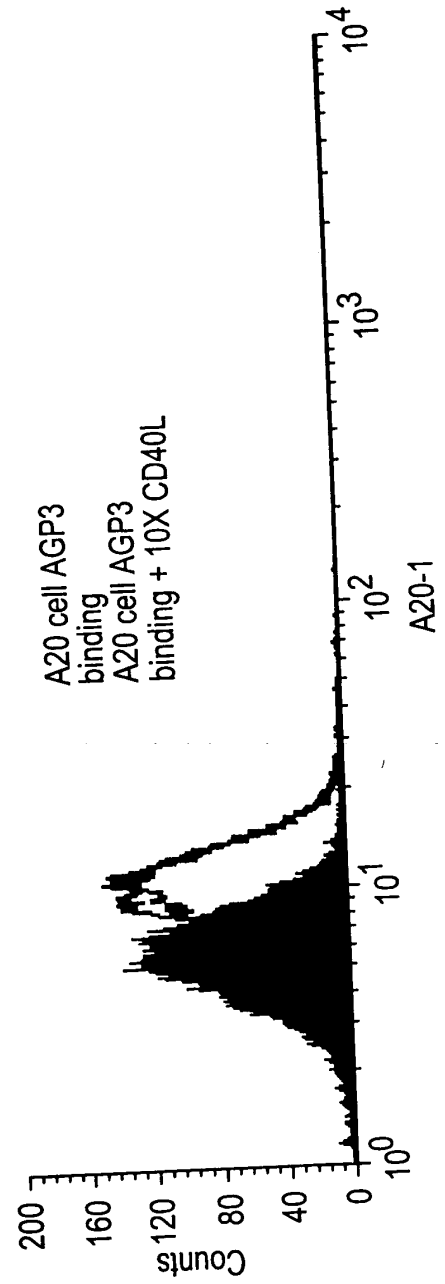


FIG. 16B





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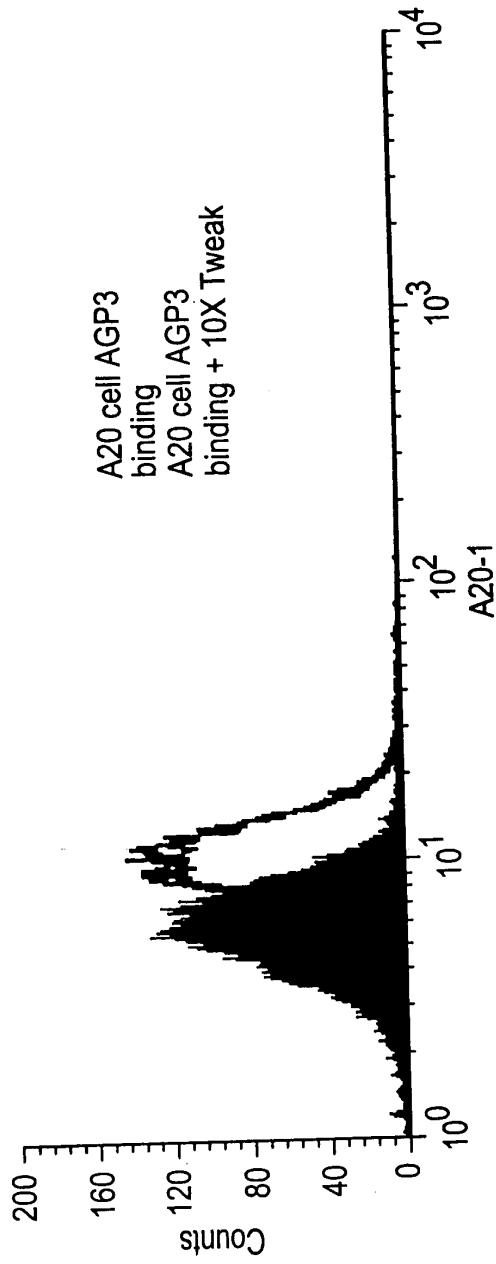


FIG. 16C

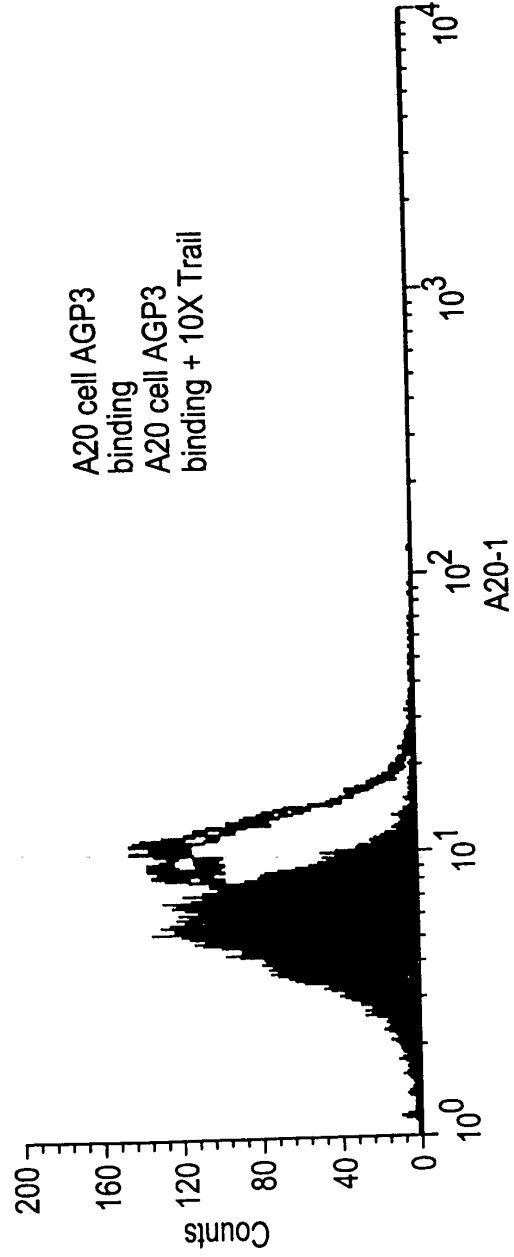


FIG. 16D



FIG. 17A

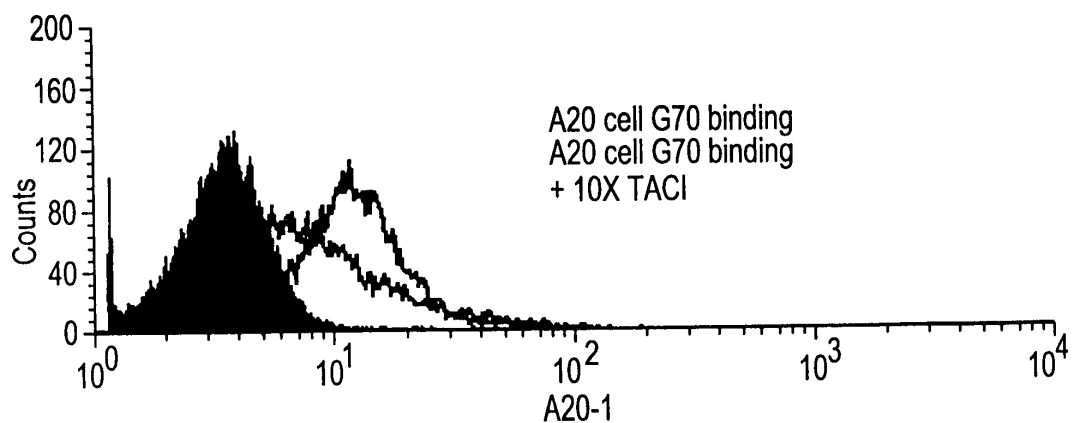


FIG. 17B

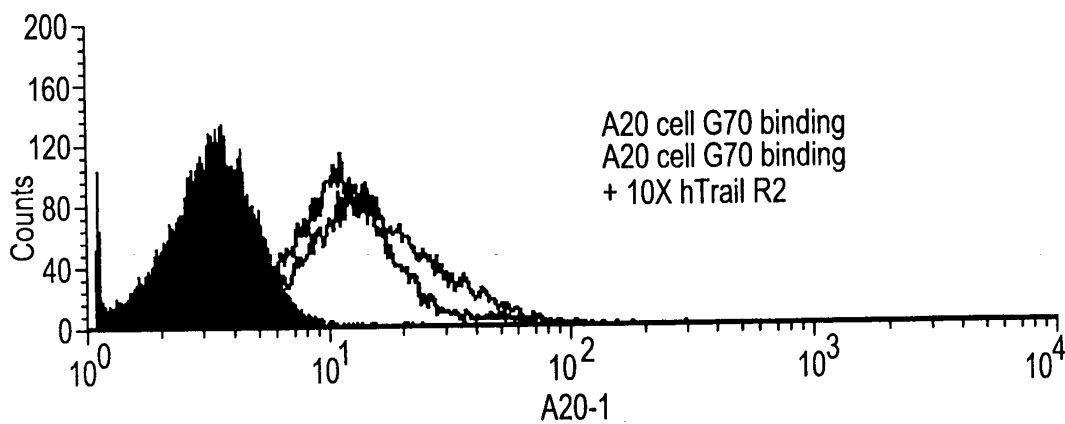
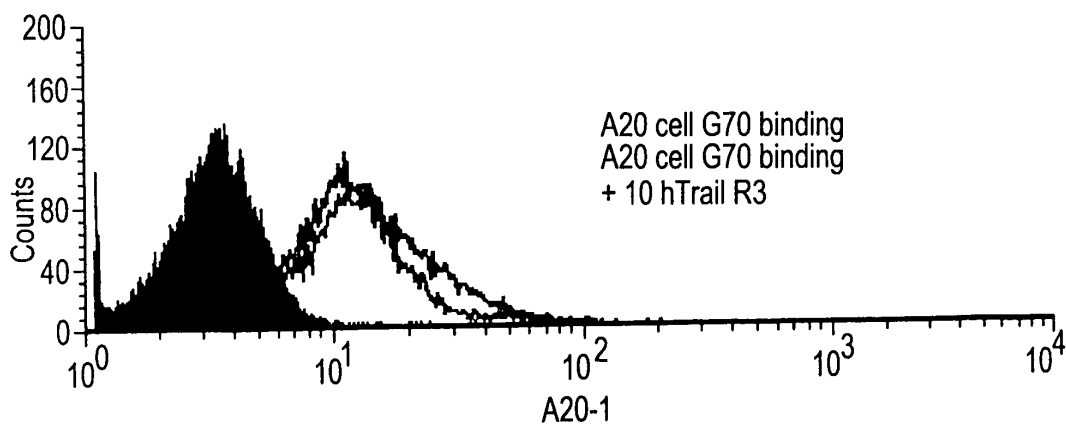


FIG. 17C



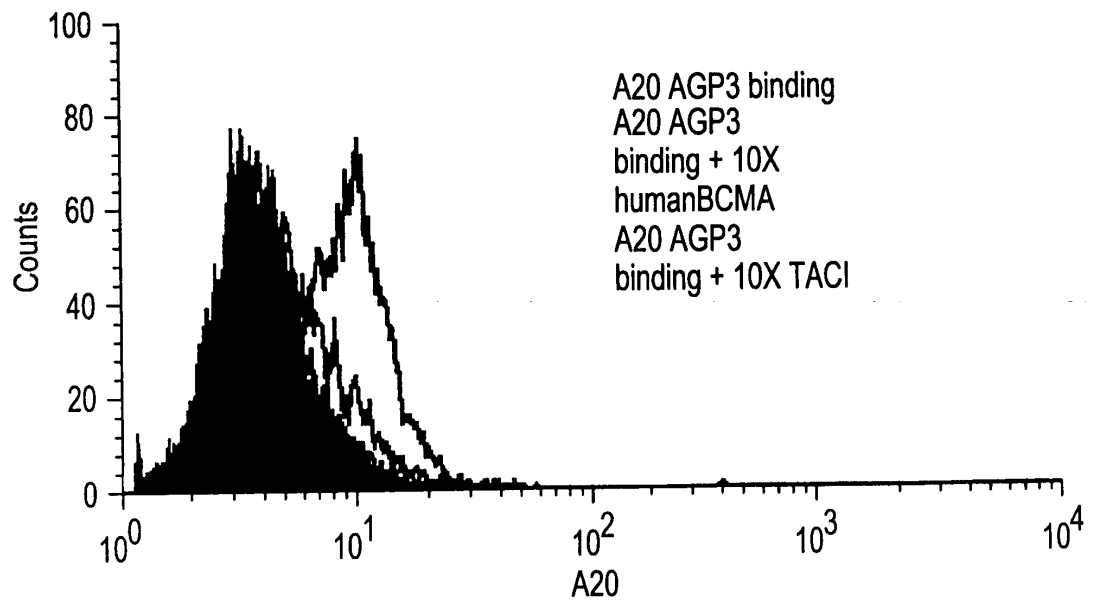
[illegible]



FIG. 19A

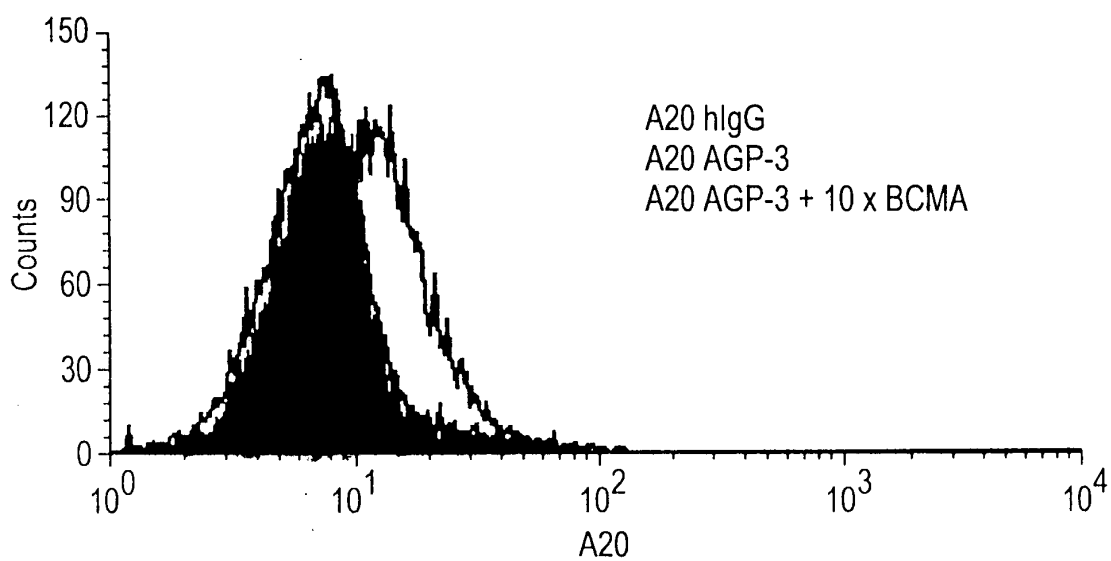


FIG. 19B

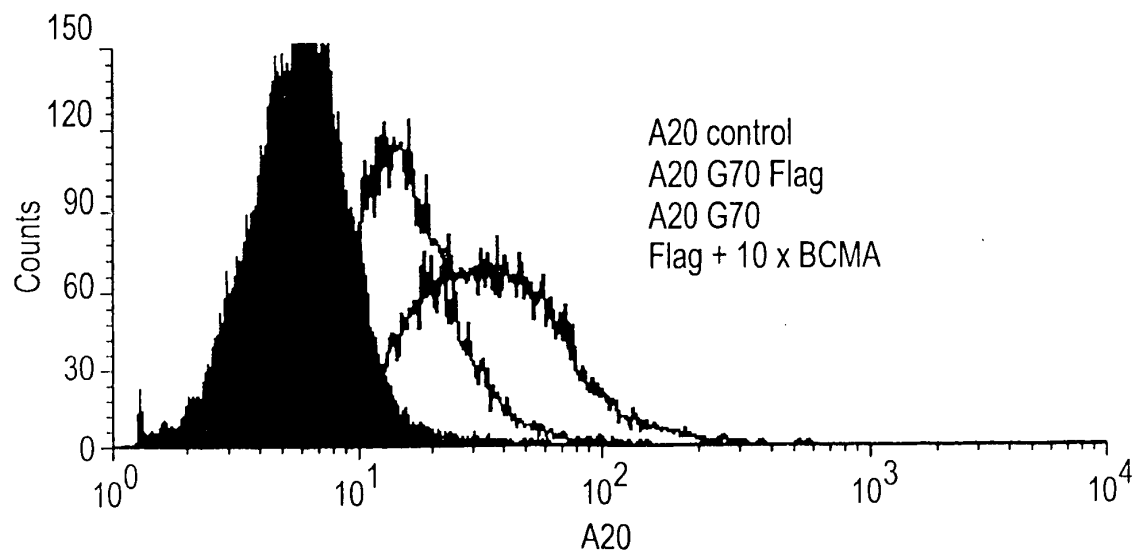


FIG. 20A

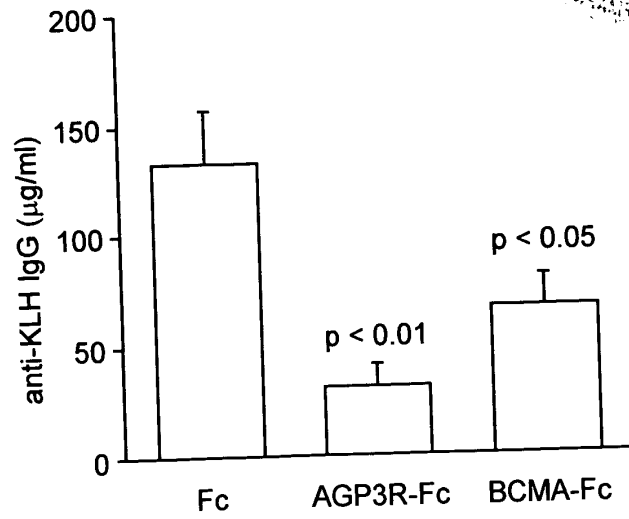


FIG. 20B

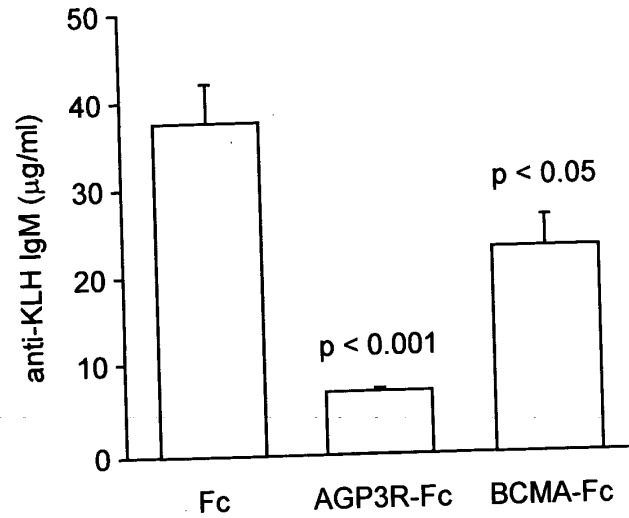
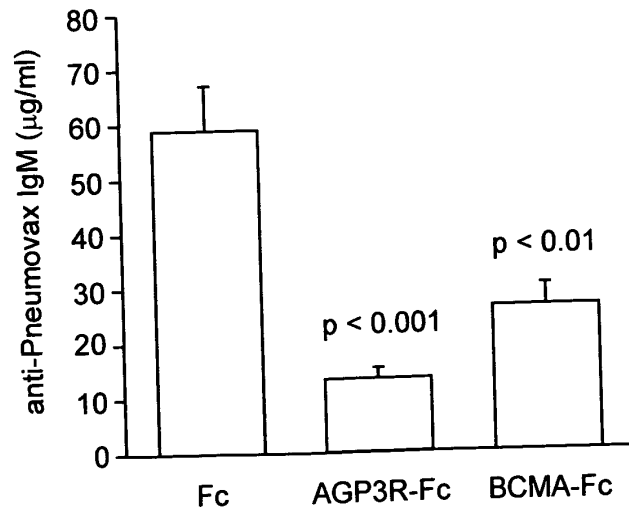


FIG. 20C



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